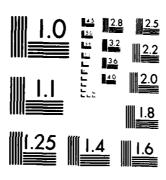
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UNITED STATES ARMY ENVIRONMENTAL HYGIENE AGENCY

ABERDEEN PROVING GROUND, MD 21010

PHASE 4

STUDY NO. 75-51-1302-84

TOXICOLOGICAL ASSESSMENT OF ABATE®

(0,0,0',0'-TETRAMETHYL-0,0'-THIO-DI-P-PHENYLENE PHOSPHOROTHIOATE)

ADMINISTERED ORALLY AND DERMALLY

TO MATED FEMALE RABBITS

APRIL 1983

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SECURITY CLASSIFICATION OF THIS PAGE (When Date Entered)

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ABATE® Cholinesterase	• •	
ABATE® Cholinesterase	Pregnancy Phenyle	-Tetramethyl-0,0'-Thio-Di-P-
ABATE® Cholinesterase Dermal Administration Gestation Oral Administration	Pregnancy Phenyle Rabbit	-Tetramethyl-0,0'-Thio-Di-P- ne Phosphorothioate



DEPARTMENT OF THE ARMY Mr. Angerhofer/or1/AUTOVON U. S. ARMY ENVIRONMENTAL HYGIENE AGENCY I

ABERDEEN PROVING GROUND, MARYLAND 21010

HSHB-OT/WP

2 No 1 1 1/4.

SUBJECT: Phase 4, Study No. 75-51-1302-84, Toxicological Assessment of ABATE (0,0,0',0'-Tetramethy)-0,0'-Thio-Di-P-Phenylene

Phosphorothicate) Administered Orally and Dermally to Mated

Female Rabbits, April 1983

Executive Secretary Armed Forces Pest Management Board Forest Glen Section, WRAMC Washington, DC 20307

EXECUTIVE SUMMARY

The purpose, essential findings, and major recommendations of the inclosed report follow:

- a. Purpose. The purpose of this study was to assess the teratologic potential of ABATE® following repeated oral or dermal administration of the compound to pregnant rabbits during the major period of organogenesis.
- b. Essential Findings. There were no teratologic effects in New Zealand White rabbits associated with repeated oral or dermal administration of ABATE at levels which produced a toxic effect.
- c. Major Recommendations. Perform chronic studies to determine any potential hazard associated with long term use of ABATE. These studies have shown that ABATE does not cause teratologic effects in laboratory rabbits.

FOR THE COMMANDER:

1 Incl as

Colonel, MGA Director, Occupational and Environmental Health

CF: HQDA (DASG-PSP) wo incl Cdr, HSC (HSPA-P) Comdt, AHS (HSHA-IPM) Dir, Advisory Ctr Div Tox, NRC (2 cy) USDA, ARS, Southern Region (3 cy) USDA, ARS, Southern Region (COL Moussa) USDA, ARS (Dr. Terrence McGovern)



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REPLY TO ATTENTION OF HSHB-OT/WP

DEPARTMENT OF THE ARMY

U.S. ARMY ENVIRONMENTAL HYGIENE AGENCY ABERDEEN PROVING GROUND, MARYLAND 21010

PHASE 4

STUDY NO. 75-51-1302-84
TOXICOLOGICAL ASSESSMENT OF ABATE®
(0,0,0',0'-TETRAMETHYL-0,0'-THIO-DI-P-PHENYLENE PHOSPHOROTHIOATE)
ADMINISTERED ORALLY AND DERMALLY
TO MATED FEMALE RABBITS
APRIL 1983

1. AUTHORITY. Letter, HSPA-H, US Army Health Services Command, 20 October 1976, subject: Investigational New Drug Application for ABATE Pediculicide, with inclosure, letter, AFPCB, Armed Forces Pest Control Board, 13 September 1976, same subject.

2. REFERENCES.

- a. Memorandum for Record, SGRD-UWF-B, Walter Reed Army Institute of Research, 18 July 1978, subject: ABATE Pediculicide.
- b. Letter, HSE-LT, this Agency, 5 December 1977, subject:
 Investigational New Drug Application for ABATE Pediculicide (Phase I).
- c. Letter, HSE-LT, this Agency, 23 April 1980, subject: Phase 2, Toxicological Assessment of ABATE (0,0,0',0'-Tetramethyl-0,0'-Thio-Di-P-Phenylene Phosphorothioate), Dermal Penetration of Radio-Labeled ABATE, Study No. 75-51-1302-80, September 1977- October 1979.
- d. Letter, HSHB-LT-T/WP, this Agency, 27 September 1983, subject: Phase 3, Study No. 75-51-1302-83, Toxicological Assessment of ABATE (0,0,0',0'-Tetramethyl-0-,0'-Thio-Di-P-Phenylene Phosphorothioate) Administered Orally to Mated and Nonmated Rabbits, April 1983.
- 3. PURPOSE. This teratologic study in rabbits is designed according to the 1966 "Guidelines for Reproduction Studies for Safety Evaluation of Drugs for Human Use" distributed by FDA. The data provided by the examination of fetuses derived from rabbits treated with ABATE during the critical period of gestation will aid in estimating the teratogenic potential, if any, of that compound. The experimental design for this study is shown in Table 1.

●ABATE is a registered tradename for American Cyanamid Co., Princeton, New Jersey 08540.

Use of trademarked names does not imply endorsement by the US Army, but is intended only to assist in identification of a specific product.

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4. SUMMARY AND CONCLUSION. Studies were conducted to evaluate the potential for ABATE to produce embryotoxic or teratogenic effects in pregnant rabbits after dermal or oral administration during the 6th through 18th day of pregnancy. The ABATE preparations and dosages used in this study were:

Dermal:

Pyrax® Powder Control
Pyrax Powder (10-percent ABATE)
Pyrax Powder (2-percent ABATE)
Technical Grade ABATE

0 mg/kg/day 163 mg/kg/day 16.3 mg/kg/day 164 mg/kg/day

Oral:

Technical Grade ABATE in 10-percent aqueous acacia

32 mg/kg/day

Intraperitoneal:

Positive Control
[6-Aminonicotinamide (6-AN)] Day 9 only

4 mg/kg

Under the conditions of the experiment, the following parameters were found to be affected:

- a. Significant decreases were found in RBC cholinesterase activity in animals receiving Pyrax powder containing 10-percent ABATE dermally (163 mg/kg/day) and technical ABATE dermally and orally (164 mg/kg/day and 32 mg/kg/day, respectively).
- b. The gestation index showed only a slight (17 and 11 percent) decrease in animals receiving technical ABATE orally and dermally, while the positive control caused a significant decrease in that index.
- c. The fertility index was lower than controls, for animals receiving 10 percent ABATE in Pyrax powder dermally. This finding was not statistically significant.
- d. Technical ABATE, at a dosage of 164 mg/kg/day, dermally, was toxic to pregnant does, causing death in 6 of 15 rabbits. This dosage was also embryotoxic, causing a decrease in total implants, decreased total alive fetuses and a significant decrease in average fetal weight when compared to fetuses from Pyrax powder controls.

[●] Pyrax is a registered trademark of R. T. Vanderbilt Company, Inc., New York, New York 10017.

- e. The positive control, 6-AN, caused a decrease in the total number of fetuses and fetuses per doe, a decrease in live fetuses and live fetuses per doe, a significant increase in resorptions and resorptions per doe, early resorptions, total malformations and malformation index. The average fetal weight was significantly decreased compared to fetuses from Pyrax powder controls.
- f. No teratologic effects were observed in fetuses from rabbit does receiving ABATE as the technical grade compound or as Pyrax powder formulations
- g. These tests indicated no teratologic hazard in New Zealand White rabbits following repeated application of ABATE at levels which produced toxic effects by both dermal and oral routes.

5. BACKGROUND.

- a. The Armed Forces Pest Management Board (AFPMB), formerly the Armed Forces Pest Control Board, is coordinating the registration of ABATE as a pediculicide with the Food and Drug Administration (FDA), since a formulation of this compound is proposed for standardization for the control of lice in military programs. Negotiations with FDA for a field test program for this preparation have indicated the need for the development of an Investigational New Drug Application (IND). The only formulation for which FDA registration is to be sought is 2-percent ABATE, 98-percent Pyrax. The target species are:
 - (1) The body louse, Pediculus humanus humanus (L).
 - (2) The head louse, Pediculus humanus cepitis (DeGeer).
 - (3) The pubic louse, Pthirus pubis (L).
- b. To assist in the development of this IND, the US Army Environmental Hygiene Agency (USAEHA) was requested to conduct a teratology study in a second mammalian species, namely the rabbit (see paragraph 1, this report).
- c. The proposed patterns of military use for 2-percent ABATE pedicult-cide encompass both group and individual treatment. The group method involves the treatment of infested individuals by operators using power-driven or manually operated equipment. It is estimated that the group application method results in the application of approximately 31 grams of formulated dust (0.62 gm active ingredient) per individual.
- d. The individual treatment method involves self-treatment. For this purpose, the dust is packaged in 2-ounce (56.7 gm) shaker cans. Instructions will indicate that the entire contents of the can be used for heavy infestations and the amount to be applied may thus be 56.7 gm of formulated powder (1.134 gm ABATE).

6. MATERIALS AND METHODS.

a. Chemicals.

(1) The experimental insecticide ABATE (0,0,0',0'-thio-di-p-phenylene phosphorothioate), CAS Number 003383-96-8, is a reddish-amber viscous liquid $(d_{25}=1.587)$, with a foul odor. It is also identified or known as Bithion, Difenthos, ENT-27165, Experimental Insecticide 52160 and Temephos. It is soluble in acetone, carbon tetrachloride, ether, ethylene dichloride and toluene. It is insoluble in hexane, methyl cyclohexane and water. The molecular weight is 466.48; its empirical formula is $C_{16}H_{20}O_6P_2S_3$ and its structural formula is shown below:

The material used in these studies was supplied by American Cyanamid Company, Agricultural Division, Princeton, New Jersey, and was contained in a labeled plastic bottle. The label contained a warning statement and the name ABATE, Technical Insecticide, Active Ingredients: Temephos [0,0' (thiodi-4, 1-phenylene) bis (0,0'-dimethyl phosphorothioate] 90 percent w/w Inert Ingredient 10 percent and lot identification number L3402 R3 6/76 WG.

- (2) Pyrax powder was received by this laboratory from Insects Affecting Man Research Laboratory, US Department of Agriculture, Agricultural Research Service, PO Box 14565, Gainesville, Florida 32604. The original source of the material was R.T. Vanderbilt Company, Inc., 230 Park Avenue, New York, New York 10017. Pyrax is their tradename for pyrophyllite, a hydrous aluminum silicate.
- (3) Acacia, U.S.P. (Gum Arabic) F.C.C., Food Grade (No. 5-0430, Lot No. 421152), was procured from J.T. Baker Chemical Co., Phillipsburg, New Jersey 08865.
- (4) 6-AN (Catalog No. A-630, Lot No. 24C0920) was purchased from Sigma Chemical Co., St Louis, Missouri.

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b. Animals. Three groups of 45 each, sexually mature, dated pregnant, female New Zealand White rabbits were purchased from Marland Breeding Farms, Inc., PO Box X, Hewitt, New Jersey 07421. The rabbits were dated from time of mating, weighed between 2.9 and 3.8 kg and were received on Day 4 of gestation. The day of mating is defined as Day 0 of gestation. The three groups were received in separate shipments at 45 day intervals. Each group of 45 rabbits was randomized and subdivided into three groups of 15 rabbits each. Animals were housed in individual cages (Porter-Mathews, 16 inches X 18 inches X 24 inches) and received laboratory diet (Rabbit Ration NIH 09, Zeigler Brothers, Inc., PO Box 95, Gardners, Pennsylvania 17321) and tap water ad libitum. The room temperature was kept at 23 ±1°C, relative humidity 45-55 percent and were maintained on a 12-hour light/dark sequence (see Table 1, Experimental Design).

c. Preparation of Solutions.

- (1) The 10-percent aqueous solutions of gum acacia were used as a vehicle control and to suspend the technical ABATE (90 mg/ml) for the oral dosing. It was also used to suspend 6-AN (100 mg/ml), the positive control for teratogenic effects.
- (2) The ABATE dust formulation under consideration is a mixture of the chemical with Pyrax powder. A 2 kg unformulated lot of Pyrax powder was received from Insects Affecting Man Research Laboratory, Gainesville, Florida, as well as 2 kg of a 10-percent (w/w) ABATE/Pyrax formulation (244.44 gm technical ABATE in 1755.56 gm pyrax powder). The ABATE dust formulation was prepared by mixing pyrax with acetone (certified A.C.S grade, Fisher Scientific) and ABATE to make a slurry. The mixture was placed under a hood to allow the acetone to evaporate. The dried material was then pulverized with mortar and pestle.
- (3) The 2 percent (w/w) ABATE/Pyrax mixture was prepared by USAEHA using acetone (A.C.S, National Stock No. 6810-00-264-8955, DSA-400-75-C-44 64, Lot 1 from ASP, Inc., Landston, Virginia 23150) as described previously, but using 22 gm of the technical grade ABATE with 1002 gm of Pyrax powder and adding sufficient acetone for the slurry.
- (4) Gravimetric analyses of the two ABATE/Pyrax formulations by the Organic Environmental Chemistry Division (OECD), USAEHA, showed the 10 percent mixture to contain 10.19-percent ABATE (OECD Log Number 5254). The 2-percent mixture contained 2.3-percent ABATE (OECD Log Number 5255).
- d. <u>Treatment Schedule</u>. Under FDA Guidelines, mated female rabbits were dosed daily from Day 6 through Day 18 of gestation. Doses were adjusted each day according to body weight. All treatments followed the daily routine of gestation Days 6-18 except for the intraperitoneal dose of 6-AN which was given on Day 9 only.
- (1) The oral dosage of 32 mg/kg was based on a preliminary range-finding study in rabbits. In that study, several concentrations of

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

TABLE 1. EXPERIMENTAL DESIGN

GROUP	TREATMENT	FORMULATION	DOSAGE	TREATMENT SCHEDULE (DAYS OF PREGNANCY)	NO. OF RABBITS
I	Exposed, oral - suspension of technical grade ABATE	90 mg ABATE/mL 10% aqueous Acacia	32 mg ABATE kg/day	6-18	15
II	<u>Control, oral</u> - solutions of aqueous 10% Acacia	10% aqueous Acacia	0.5 mL/kg/day	6-18	15
111	<u>Positive Control</u> - Intra- peritoneal administration of 6-AN	100 mg 6-AN/mL 10% aqueous Acacta	2.5 mg 6-AN/kg	Day 9	15
IV	<u>Control Dust - dermal</u> application untreated Pyrax		1.81 gm Pyrax/ kg/day	6-18	15
٧	Exposed Dust - dermal application 10% ABATE in Pyrax	100 mg ABATE/ gm Pyrax	163 mg ABATE/kg/day	6-18	15
VI	Exposed Dermal application technical grade ABATE	·	164 mg ABATE/kg/day	6-18	15
VII	<u>Exposed, oral</u> - suspension of technical grade ABATE	90 mg ABATE/mL 10% aqueous Acacia	32 mg ABATE/kg/day	6-18	15
VIII	<u>Positive Control</u> - <u>Intra-</u> <u>peritoneal</u> administration of (6-AN)	100 mg 6AN/mL 10% aqueous Acacia	4.0 mg 6-AN/kg	Day 9	15
IX	Exposed Dust - dermal application 2% ABATE in Pyrax	20 mg ABATE/gm Pyrax	16.3 mg ABATE/kg/day	6-18	15

ABATE in 10-percent gum acacia were given daily by the oral route to determine a dosage which would lower plasma and RBC cholinesterase levels to about 50 percent of the pretreatment levels.

- (2) The 10-percent ABATE/Pyrax formulation and the dermal application of technical grade compound were treatments at dosages which were approximately 10 times the projected single, manual human use treatment on a mg per kg basis.
- (3) The 2-percent ABATE/Pyrax formulation approximated the projected single, manual, human-use treatment.
- (4) The dosage for 6-AN was based on an extrapolation from data in an article by Schardein, et all.

e. <u>Test Procedure</u>.

- (1) Mated animals were received on Day 4 of gestation. Treatment was initiated on Day 6 and continued through Day 18 of gestation for all groups except those receiving the single intraperitoneal dosage of 6-AN on Day 9.
- (2) Daily observations for toxicologic signs were made. All rabbits were weighed on Day 4; Days 6-18; Days 22, 25, 28 and 30. Animals were bled from the the central ear artery on Days 5, 7, 19 and 30 of gestation for determination of plasma and erythrocyte (RBC) cholinesterase activity. These clinical chemistry analyses were performed according to the method described by Garry and Routh². All animals were sacrified on Day 30 of gestation by means of an intravenous overdose of barbiturate. At this time, the brains were removed from two fetuses from each litter for the determination of brain cholinesterase activity. The brains from all does were also analyzed for cholinesterase activity.
- (3) The examination of fetuses for malformations was conducted according to the method of Wilson and Warkany 3 . The post mortem for each doe consisted of counting the conceptuses: number, location, living, dead, early resorption and late resorption. All fetuses were tagged for identification, weighed, measured and examined for external defects. Approximately one-third (1/3) of the fetuses were fixed in Bouin's fluid and examined by the Wilson technique for neural and visceral defects. The remaining two-thirds (2/3) of the fetuses were placed in 95 percent ethyl alcohol, cleared and their skeletons stained with alizarin red S and examined for the presence of anomalies 5 .

f. Evaluation of Data.

(1) Definition of Terms. The following indices were calculated. Results appear in Appendices A-H.

Index of fertility:	pregnant animals total number of mated animals	X 100
Index of viable births:	alive normal fetuses total number of fetuses	x 100
Index of dead births:	dead normal fetuses total number of fetuses	X 100
Index of resorptions:	total number of resorptions total number of implantations	X 100
Index of variations:	total number of variations total number of fetuses	X 100
Index of malformations:	total number of malformations total number of fetuses	X 100
Index of gestation:	total number of litters total number of females pregnant	X 100

Variations: All runts and anomalies.
Early resorptions: Placental remains only.
Late resorptions: Placental and fetal remnants.
Runts: A fetus weighing 70 percent or less of the average weight of its litter.

(2) Statistical Analysis. Applicable fetal parameters, cholinesterase values and body weights were analyzed statistically using the Student's "t" test with significance selected at the 0.05 level of probability.

7. RESULTS.

a. Exclusion of Test Groups. Data from rabbit groups I, II, and III are considered inadequate and are not reported or discussed. Dosing methods resulted in a high percentage of maternal deaths in groups I (11/15) and II (13/15) by the end of the study. This high mortality rate resulted in an insufficient number of animals reaching term and it was decided that such a sample size was too small upon which to base any conclusions. Similarly, the positive control Group III received a 0.3 mg/kg dosage of 6-AN, intraperitoneally, on Day 9 of gestation instead of 3.0 mg/kg. Data from this group was also considered incomplete and invalid and was not included in the body of the study.

- b. The following results are given for Groups IV through IX:
 - (1) Maternal Parameters.
- (a) Clinical Picture of Females. Animals of all groups were received in good condition and showed a smooth, shiny hair coat. A few had diarrhea for 2-3 days after arrival, but this condition soon disappeared and the fecal pellets appeared normal for the duration of the test period.
- (b) Weight Gain of Females. Body weights remained essentially, unchanged or showed slight gain during the treatment period (Days 6-18 of gestation). These data are summarized in Appendix I. Individual data are presented in Appendices J-P.
- (c) Toxicity. Repeated treatment of rabbits with dermal or oral administration of technical grade ABATE caused toxic responses with deaths accuring in each group of animals. No skin irritation resulted from dermal application of technical grade ABATE or the Pyrax formulations.
- (d) Cholinesterase Activity. Oral and dermal application of technical grade ABATE and Pyrax with 10-percent ABATE caused a significant reduction in RBC cholinesterase activity after 13 days of treatment. This activity remained depressed in these groups at sacrifice. Maternal brain cholinesterase activity was not affected in any treatment group. Plasma cholinesterase activity was depressed after 13 days of treatment with dermal application of technical grade ABATE and Pyrax with 10-percent ABATE. The plasma cholinesterase activity following oral ABATE treatment indicated a decreasing trend, but was not statistically significant (p <.05). Plasma cholinesterase activity in dermal technical ABATE rabbits remained depressed at sacrifice. Plasma and RBC cholinesterase activities were not depressed in the 2-percent ABATE pyrax group. Cholinesterase data are presented in Appendices Q-FF.
- (e) Gestation and Fertility Indices. The gestation index was significantly reduced in the 6-AN group and was reduced by 11 and 17 percent in the dermal and oral technical grade ABATE groups respectively. The fertility index was reduced in the group receiving dermally applied Pyrax with 10-percent ABATE.
- (f) Necropsy Findings. No gross changes in tissues or organs of the female rabbits were found at necropsy.
 - (2) Fetal Parameters.
- (a) Index of Dead and Live-born Fetuses. No differences of biological relevance were found upon comparison of the groups.
- (b) Implantations. Implantations per doe were 21 percent lower than Pyrax controls in does receiving technical grade ABATE dermally. This finding is not considered to be compound related since implantation would have occured prior to the start of treatment.

- Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83
- (c) Index of Resorptions. Resorption sites were found in does from all treatment groups. As expected, the positive control group had a much greater number of resorptions than test or negative control groups.
- (d) Abnormalities/Anomalies. All true malformations (soft tissue or skeletal defects such as gastroschisis, exencephaly, cleft palate) are classified as abnormalities. Anomelies are considered to be minor varients from the normal, such as unossified sternebrae and retarded ossification of the fontanella. A single malformed fetus from each of Groups IV, V and VII is considered to be a natural occurance and not related to the test compound. The number of varients showed no compound related differences. Group VIII, given the known teratogen 6-AN, showed an expected high incidence of malformations. A summary of fetal examination findings is presented in Appendix E.
- (e) Average Number of Fetuses. Group VIII (6-AN) showed a significant decrease in the absolute total of fetuses and the mean number of fetuses per doe. Group VI (dermal technical ABATE) showed some reduction in the total and mean number of fetuses, but this is considered to be an expression of the maternal and fetal toxicity of dermally applied technical grade ABATE.
- (f) Average Fetal Weight. Significant differences were found between control and test fetuses from the dermal technical ABATE and 6-AN groups. No differences between control and the remaining treatment groups occurred at p <.05. Data on fetal size is presented by litter in Appendix GG.

RICHARD A. ANGERHOFER

Richard Ca Cazalot

Chief, Toxicology Division

Biologist

Toxicology Division

MAURICE H. WEEKS

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APPENDIX A
SUMMARY OF MATERNAL AND FETAL PARAMETERS

	Dermal Pyrax Control IV	Dermal Pyrax 10: ABATE V	Dermal Technical ABATE VI	Oral ABATE VII	I.P6-AN Positive Control VIII	Dermal Pyrax 2% ABATE IX
Females mated	13	14	8	10	14	12
Females pregnant	10	9	6	9	11	10
index of Fertility (%)	77	64	75	90	79	83
Litters	10	9	5	8	5	10
Index of Gestation (%)	100	100	83	89	45*	100
Implantations total	94	96	47	89	99	113
Implantations per doe	9. 40	10.67	7.83	9. 89	9. 00	11.30
Fetuses, total	87	82	33	65	16	92
Fetuses per doe	8.70	9.11	5. 50	7.22	1.45*	9. 20
Alive fetuses, total	86	81	33	65	16	92
Alive fetuses per doe	8. 60	9.00	5. 50	7.22	1.45*	9. 20
index of viable fetuses (%)	98.85	97.59	100	100	100	100
Dead fetuses, total	1	1	0	0	0	0
Dead fetuses per doe	0. 10	0.11	0	0	0	0
Index of deed births (%)	1.15	1. 23	0	0	0	0
Resorptions, total	7	14	14	24	83	21
Resorptions per doe	0.70	1.56	2. 33	2.67	7.55*	2. 10
ndex of Resorptions (%)	7.45	14.58	29. 79	26.97	83.84°	18. 58
Early resorptions	4	10	14	24	83	20
Late resorptions	3	4	0	0	0	1
Veriants, total	4	2	2	3	1	0
Variants per doe	0.40	0. 22	0. 33	0. 33	0.09	0
index of verietions (%)	4.60	2.41	6.06	4.62	6. 25	0
felformetions, total	1	. 3	0	1	53	0
index of walformations (%)	1. 15	3. 61	0	1.54	331*	0
lunts	2	1	2	1	1	0
lverage fetus weight (g)	48. 20	48. 60	40.80*	49.70	37.60*	52.40

^{*} Significantly different from dermal pyrax control at .05 level of probability.

APPENDIX B EXAMINATION OF SKELETAL AND SOFT TISSUE STRUCTURES FOR MALFORMATIONS AND VARIATIONS

Animai No.	Observations	Fetus Quantity	Quality
	Group IV (Control)		
48	talipes equinovarus	1	
52	parietal bones overlap frontals	1	٧
	Group V (Dermai Pyrax 10% ABATE)		
90	ancephalia	1	
	thoracoschisis	ì	
	gastroschisis	ī	
	Group VII (Oral technical ABATE)		
94	enlarged fontanella	1	٧
95	parietal bones overlap frontals	i	Š
105	tibias, fibulas and femurs not	•	•
103	completely formed	1	
	Group VIII (IP-6-AN)		
107	spina bifida	1	_
107	microopthalmia		
	cleft palate	3	
	talipes equinovarus	် င	
	qastroschisis	5 3 5 1 2 3 3	
	fused ribs	į	m
113	microopthalmia	2	m
113		્ર	
	talipes equinovarus	3	
	gastroschisis	1	
	fused ribs	Ş	
114	spina bifida	1.	
	microopthalmia	Š,	•
	cleft palate	1	
	cranjoschisis	1	#
~ .	gastroschisis	1	•
	webbed feet	1	-
	fused ribs	1	
118	microopthalmia ,	3	
	cleft palate	2	
	talipes equinovarus	<u>1</u>	
120	microopthalmia	3	
	cleft palate	3	
	cranioschisis	1	
	hydrocephalia	1	
	talipes equinovarus	2	
	fused ribs	1	
	frontal & parietal bones		
	deformed	2	

⁼ variation
= malformation

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APPENDIX C

INDIVIDUAL MATERNAL AND FETAL PARAMETERS GROUP IV DERMAL PYRAX CONTROL

No.	Mated	Pregnant	Resorptions Pregnant Implantations Early Late	Resorptions Total No. Early Late of Fetus	Total No. of Fetuses		Al ive N	Dead Alive Malformations
46	+	•						
47	+	Died before term	e term					
48	+	+		0	6	•	00	-
49	+	+	13	1 0	12	0	12	. 0
20	+	+	_	0	9	· C	יב	
51	+	•	•	•	•	•)	•
25	+	+	σ	0	6	0	σ	c
53	+	1		•	•	•	•	•
54	+	•		0	11	0	11	c
22	+	+	∞	1 0	7	0	-	· C
26	+	+	75	0	12	· c	12	· c
27	+	+	_	2 0		• 0	י יי	
28	+	+	10	0	10	0	10	0
29	+	+	· o	0	9	0	9	0

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APPENDIX D
INDIVIDUAL MATERNAL AND FETAL PARAMETERS
GROUP V
DERMAL PYRAX WITH 10 PERCENT ABATE

Ko.	Mated	Pregnant	Pregnant Implantations	Resorptions Early Late	Late	Resorptions Total No. Early Late of Fetuses	Dead	- 1	Alive Malformations
61	+	+	16	0	0	16	0	16	0
29	+	+	9	0	0	g	0	9	0
63	+	ı					,	1	•
64	+	+	12	-		임	0	10	0
9	+	+	യ	0	0	ထ	0	œ	0
99	+	+	13	~	2	O	0	O	0
29	+	+	10	0	-	σ	0	o	0
89	+	+	Aborted						·
69	+	+	10	0	0	10	0	10	0
2	+	•)) 	1
7	+	1							
72	+	+	11	7	0	4	c	4	C
73	+	1	1		,	•	•	•	•
74	+								
90	+	+	10	c	C	10		σ	~

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APPENDIX E INDIVIDUAL MATERNAL AND FETAL PARAMETERS GROUP VI DERMAL TECHNICAL ABATE

Animal No.	Mated	Pregnant	Implantations	Resorptio Early La	Resorptions Total No. Pregnant Implantations Early Late of Fetuses Dead Alive Malformations	Dead	Al ive	Malformations
75	•	•	α	c	α	c	α	•
29	• +	. 1	•)	•	•	,	
11	+	+	∞	4	4	0	4	•
82	+	Died before term	e term					
79	+	Died before term	e term					
8	+	Aborted	•					
81	+	+	~	∞	0	0	0	•
85	+	•						
83	+	Died before term	e term					
84	+	+	~	0	8	•	∞	•
82	+	Sacrificed	Sacrificed before term (moribund)	oribund)				
86	+	+	9	~	4	0	4	•
87	+	Died before term	e term					
88	+	Died before term	e term					
88	+	+	6	0	6 0	0	O	•

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APPENDIX G
INDIVIDUAL MATERNAL AND FETAL PARAMETERS
GROUP VIII
INTRAPERITONEAL 6-AN

No. Mated Pregnant Implantations Early Late of Fetuses Dead Allv 106 + bied before term 5 0 5 0 5 107 + + 10 5 0 0 0 0 108 + + 11 11 0	Animal				Resorp	tions	Resorptions Total No.			
+ Died before term	2	Mated	- {	Implantations	Early	Late	of Fetuses	Dead	Alive N	Alive Malformations
+ + + + 10	106	+	Died before	term						
+ + + + 23 - 23 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	107	+	+	01	S	0	ഗ	0	S	17
+ + + + + 5	108	+	+	11	Ξ	0	0	0	0	0
+ + + + + + + + + + + + + + + + + + +	109	+	•							
+ + + + + + + + + + + + + + + + + + +	110	+	+	23.	23	0	0	0	0	0
+ + + + + + + + + + + + + + + + + + +	111	+	+	12	12	0	0	0	0	0
+ + + + + + + + + + + + + + + + + + +	112	+	•							
+ + + + + + + + + + + + + + + + + + +	113	+	+	ဖ	m	0	m	0	ო	6
+ + + + + + + + + + + + + + + + + + +	114	+	+	ß	က	0	7	0	2	œ
+ + + + 7 4 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	115	+	+	S	വ	0	0	0	0	0
+ + + 7 7 7 0 0 0 0 0 0 0 0 0 0 0 0 0 0	116	+	+	4	4	0	0	0	0	0
+ + + 7 4 0 3 0 0 + + + + + 9 6 0 3 0	117	+	•					ı	ı	•
+ + + + 9 6 0 3 0 0 3 0 0 3 0 0 0 0 0 0 0 0 0 0 0	118	+	+	7	4	0	ო	0	m	ဖ
0 9 6 + +	119	+	+	7	7	0	0	0	.0	0
	120	+	+	O	છ	0	က	0	က	13

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APPENDIX F

INDIVIDUAL MATERNAL AND FETAL PARAMETERS GROUP VII ORAL TECHNICAL ABATE

No. Mated Pregnant Implantations Early Late of Fetuses 91	Animal				Resorptions	ions	Total No.			
+ + + + 9 0 0 0 1	No.	Mated		Implantations	Early	Late	of Fetuses		Alive M	Dead Alive Malformations
+ Died before term + + + 10 0 0 1 + + 11 3 0 + + 11 3 0 + Died before term + + + 12 12 0 + + + 9 3 0 0 + + + + + 9 0 0 0 + + + + + + 9 0 0 0 + + + + + + + 9 0 0 - **A **In the state of the st	91	+	+	თ	0	0	o	0	6	0
+ + + + 10 0 0 1 + + + + 11 3 0 1 + Died before term + + + + + + + + + + + + + + + + + + +	35	+	Died befor	e term						
+ + + + 11 3 0 + + + + 10 3 0 + Died before term + Died before term + + + 12 12 0 + Died before term + Died before term + + + 9 3 0 + + + + 9 0 0	93	+	+	10	0	0	10	0	10	0
+ + + 11 3 0 + Died before term + Died before term + + + 12 12 0 + Died before term + Died before term + + + 9 3 0 + + + 9 0 0	94	+	+	11	က	0	ထ	0	ω	0
+ + + + 10 3 0 + Died before term + Died before term + Died before term + Died before term + + + + 9 3 0 + + + + 9 0 0	95	+	+	11	က	0	&	0	ω	0
+ Died before term + + + 9 3 0 + + + 9 0 0	96	+	+	• 10	က	0	7	0	7	0
+ Died before term +	6	+	Died befor	e term						
+ + + 12 12 0 + Died before term + Died before term + + + 8 0 0 + + + + 9 0 0	88	+	Died befor	e term						
+ + + 12 12 0 + Died before term + Died before term + + + 8 0 0 + + + 9 0 0	66	+	•							
+ Died before term + Died before term + + + 8 0 0 + + + 9 0 0	100	+	+	12	12	0	0	0	0	0
+ Died before term + + + 9 3 0 + + + 8 0 0 + + + 9 0 0	101	+	Died befor	e term						
0 0 0 0 6 + + + + + + + + + + + + + + +	102	+	Died befor	e term						
0 0 0 8 + + + +	103	+	+	6	ო	0	9	0	9	•
0 0 6 + +	104	+	+	œ	0	0	ω	0	Φ	•
	105	+	+	6	0	0	თ	0	თ	

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APPENDIX H
INDIVIDUAL MATERNAL AND FETAL PARAMETERS
GROUP IX
DERMAL PYRAX WITH 2 PERCENT ABATE

8	Mated	Pregnant	Resorption Prequent Implantations Early Late	Kesorp Early La	tions ate	Resorptions lotal No. arly Late of Fetuses	Dead	Alive 1	Alive Malformations
121		+	10	-	6	6	0	On.	•
122	+	+		. –	0	φ.	0	ဖ	1
123	+	+	. On	~	0	7	0	~	1
124	+	+	15	4	0	11	0	=	•
125	+	•							
126	+	+		2	0	6	0	O	•
127	+	+	2	7	0	œ	0	œ	•
128	+	+	11	7	0	6	0	6	•
129	+	Died before term	e term						
130	+	+		-	0	21	0	ន	•
131	+	+	16	4	-	#	0	11	•
132	+	+	13	-	0	12	0	12	•
133	+	Died before term	e term						
134	+	Died before term	e term						
135	+	•							

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APPENDIX I

MEAN BODY WEIGHTS (kg) - Female Rabbits

			Day of	f Gestatio	n	
Group		Day 4 Day	Day 6 Treatment	Day 9	Day 18 Final	Day 30
Treatment		Received	Starts		Treatment	Sacrifice
Dermal IV	ž	3.91	3.98	3.92	3.97	4.10
Pyrax Control	SD	.38	.46	.38	.41	.41
Dermal V	ž	3.74	3.68	3.68	3.69	3.90
Pyrax 10% ABATE	SD	.38	.47	.50	.47	.53
Dermal VI	x	3.77	3.71	3.71	3.69	3.64
Tech ABATE	SD	.31	.28	.33	•44	.47
Oral VII	x	3.63	3.56	3.55	3.70	4.1
ABATE	SD	.24	.26	.27	.51	.16
IP-6AN VIII	X.	3.53	3.57	3.63	3.47	3.76
Positive Control	SD	.24	.27	.31	.37	.33
Dermal IX	ž	3.59	3.57	3.46	3.52	3.92
Pyrax 2% ABATE	SD	.26	.20	.18	.36	.53

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APPENDIX J

STATISTICAL ANALYSES OF BODY WEIGHT CHANGES (Percent body weight changes) - Analysis by Student "t"

	Day 30/Day 4 Complete			Day 18/Day 6 Treatment	DAY.	<u>+</u>	DAYS OF GESTALION Day 30/Day 18 Post treatment			Day 6/Day 4 Preexposure	
Group/Treatment	Mean & Change (±SD)	5	44	Mean & Change (±SD)	늄	44	Mean % Change (±SD)	P	4	Mean \$ Change (±SD)	8
IV Dermal Pyrax Control	105	•	,	100	•	1	102	•	•	102	•
	7	٠.		9	•	•	9		•	s	٠
V Dermal Pyrax 10% ABATE	104	56 0	0.19	99	27	0.07	106	92	1.25	*86	82
VI Dermal Technical ABATE	96.	19 1	1.49	66	52	0.29	26	19	1.23	î 66	58
***	15		. 3	φ ;	' ;		12	' ;		us (• 9
VII OFAI ABAIL	1111* 6	7 7	2.20	103 12	57 '	66.0	106 4	77	i. 66	8 8 8	27
VIII I.P. 6-AN (Positive Control)	108	25 0	0.92	88 9	97	1.07	108* 5	25	2.49	101 6	≈ '
IX Dermal Pyrax 2% ABATE	107	23 0	0.51	98 10	- 52	0.60	108	23	1.87	100	88 '

1.87

0.30

1.12

* Significant differences (p<.05) of body weight changes (%) comparing pyrax control with each of the various treatments.

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	WEIGHTS(kg)	108
×	BODY 1V	CONT
APPENDIX K	MATERNAL BOI	DEPMA! DYRAX CONTROL
	INDIVIDUAL MATERNAL BODY WEIGHTS(kg)	משט

							j	DAY OF (GESTATION	NO									
Animal Number	Day 4	o S	Day 6 Treat- ment Starts	Day 7	Day 8	Day o	Day 10	Day 11	Day 12	Day 13	Day 14	Day 15	Day 16	Day 17	Day 18 Final Freat- ment	Day 19	23 23	25 25	30 30 30
46	6.3	4.45	4.37	4.40	4.26	4.36	4.34	4.40	4.42	4.44	4,49	4.56	4.52	4.43	4.43	4.41	4.63	4.71	4.63
4 4 8	2. 4. 2. 6.	4,75	4.70	3.38 7.72	2.83 2.66	4,69	4.69 69	4.77	4.72	4.81	4.80	4.81	4.82	4.81	4.79	4.75	4.79	4.83	4.93
6	4.6	4.62	4.53	4.45	4.40	4.50	4.49	4.49	4.51	4.58	4.61	4.57	4.62	4.52	4.53	4.59	4.73	4.69	4.43
20	3.5	3,49	3.41	3.44	3,39	3.60	3.46	3.44	3,46	3.57	3.56	3,67	3.61	3.66	3.65	3.65	3.89	3.96	3.97
51	8.8	3,91	3.75	3.72	3.69	3.68	3.58	3.61	3,63	3.51	3,42	3,47	3.41	3.44	3.49	3.51	3.75	3.74	DIED
52	3,8	3.78	3.52	3.55	3.51	3.61	3.57	3.57	3.72	3.72	3.71	3.76	3.86	3.79	3.74	3.74	3.84	3.83	3.86
53	3.3	3.39	3,45	3,41	3,73	3.44	3.45	3.41	3,48	3.42	3,41	3,45	3.53	3,43	3.44	3.49	3.54	3.44	3.57
55	3.6	3,55	3.59	3.57	3.52	3.50	3.52	3.54	3.50	3.52	3.62	3.66	3.67	3.60	3.65	3.62	3.71	3.78	3.89
55	3.8	3.65	3.87	3.80	3.82	3.81	3.73	3.85	3.83	3.78	3.89	3,95	3.99	3.92	3.97	3,85	4.12	4.23	4,34
95	3.8	3.70	3.98	4.07	3.94	3.93	3.88	3.90	3,92	3,95	4.05	4.06	4.06	3.99	4.06	4.12	4.30	4.29	3,54
57	4.2	3.90	4.60	4.06	4.05	4.18	4.01	4.01	4.02	4.02	4.04	4.01	4.00	3.95	3.92	3.86	3.70	3,75	3.86
58	4.2	4.00	4.70	4.31	4.23	4.29	4.07	3,83	4.25	4.30	4.41	4.43	4.45	4.34	4.40	4.40	4.39	4.38	4.32
29	3.6	3.50	3.67	3.50	3.66	3.65	4.05	3.64	3.66	3.71	3.74	3.69	3.70	3.76	3.73	3.72	3.95	4.01	3.93
8	3.7	3.50	3.70	3.50	3.73	3.93	4.05	4.15	3.86	3.86	3.98	3,90	3.89	3.90	3.80	3.87	3.97	3.95	3,99
S + ×	3.9	3.87	3.98 0.46	3.89	3.8 9 0.36	3.92	3.90	3.87	3.89 0.40	3.94	3.98	4.00	4.01	3.97	3.97	3.97	4.09	4.11	4.10

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APPENDIX L

INDIVIDUAL MATERNAL BODY WEIGHTS (kg) GROUP V DERMAL PYRAX WITH 10 PERCENT ABATE

	30 30	84.4444444.64.6.84 64.4444.64.6.84.8.84.4.6.84.8.84.8	3.90
	Day 25	64.0.44.44.0.64.0.64.0.94.0.94.0.94.0.94	3.89
	Day 23	44.116 44.116 44.116 43.128 43	3.83
	Day 19	33.94 33.98 33.98 33.35 33.35 43.08 33.57 43.08 43.08	3.70
	Day 18 Final Treat- ment	6.00 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	3.69
	Day 17	65 8 8 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9	3.67
	Day 16	3.53 3.53 3.53 3.53 3.53 4.53 4.53 4.53	3.68
	Day 15	3.74 3.74 3.92 3.92 3.92 3.92 3.32 3.32 4.60 4.60	3.71
	Day 14	3.51 2.65 3.77 3.26 3.86 3.80 3.20 3.20 3.55 3.35 3.35 4.88	3.71 0.51
TION	Day 13	3.48 3.751 2.70 2.70 3.86 3.83 4.35 3.42 3.75 3.75 3.77 3.61	3.69 0.51
GESTATION	Day 12	3.743 2.63 2.63 2.63 3.39 3.21 3.21 3.59 4.48	3.67
DAY OF	Day 11	2.51 2.52 2.53 2.53 2.53 2.53 2.53 2.54 2.54 2.55 2.55 2.55 2.55 2.55 2.55	3.70
	Day 10	8 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2	3.66
	Day 9	33.788 33.673 33.653 33.743 33.742 33.742 5133 5133	3.68
	Day 8	3.50 3.50 3.50 3.50 3.50 3.50 3.50 4.17 5.50	3.67
	Day 7	33.62 33.65 33.05 33.05 33.05 33.71 33.72 33.73 52 52	3.69
	Day 6 Treat- ment Starts	50000000000000000000000000000000000000	3.68
	Day 5	6.6.6.6.4.4.6.6.6.6.4.4.6.6.6.6.6.4.4.6	3.75
	Day 4	พนพลนลลลพนพพพนพน พับพัฒชังสลักษ์สลักลังกับ	3.7
}	Anfma] Number	65 65 65 65 65 65 65 65 65 65 65 65 65 6	S I+ ×I

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INDIVIDUAL MATERNAL BODY WEIGHTS (kg) GROUP VI DERMAL TECHNICAL ABATE

APPENDIX M

		30	3.73 3.58 3.58	2.86 3.51 4.12	3.83 ED 4 A	4.29 3.64 0.47
		08y 25	3.60 4.05 3.53 Apr	3.57	3.90 SACRIFIC 3.15	11 Mar 4.10 3.70 0.30
ļ	i	0ay 23	3.60 3.40 01ED 3	3.65 3.55 3.80	3.90 3.75 3.10	3.68 0.36
		Dag 19	3.55 4.65 3.60 2.80 3.25	4.00 3.65 3.70	3.99 3.70 3.10	3.25 4.00 3.63 0.48
i		Day 18 18 Final Treat- ment	3.65 4.70 3.80 2.85 3.55	3.70 3.85	3.80 3.70 3.15	3.35
		0ay 17	3.60 3.90 3.90 3.65	4.15 3.65 3.85	3.85 3.80 3.10 28 Mar	3.45 3.95 0.45
	ļ	Day 16	2.95 2.95 3.95	4.15 3.65 3.80	3.95 3.90 3.05 DIED	3.55 3.95 3.75 0.47
		Day 15	3.55 3.90 3.05	4.05 3.70 3.80	3.10	3.60 3.90 0.43
		0ay 14	3.50 3.90 3.20 3.70	3.60 3.90	3.95 4.05 3.10	3.55 3.90 3.73 0.41
	No	Day 13	3.95 3.35 3.35	4.15 3.55 3.90	3.90 3.85 3.35	3.45 3.85 3.72 0.42
	GESTATION	Day 12	3.75 3.90 3.35 3.65	4.10 3.50 3.85	3.90 2.95 3.45	3.55 3.80 3.74 0.38
	P.	Day 11	3.85 3.35 3.35	3.60 3.80	3.92 3.92 3.45	3.55 3.80 3.73 0.32
	DA	Day 10	3.85 3.95 3.30 3.70	3.50 3.90	3.95 3.10 3.50	3.50 3.60 3.73 0.32
		Day 9	3.93 3.93 3.00	3.55 3.75 Mar	3.90 9.90 9.00 9.00	3.55 3.60 3.71 0.33
		Day 8	3.85 3.90 3.20 3.60	3.60 3.70 DIED 20	3.70 2.95 3.75	3.60 3.70 3.70 0.33
		Day 7	3.80 3.90 3.15	4 u u u 6 6 6 6	3.70 3.80 3.00 3.75	3.55 3.70 3.66 0.32
		Day 6 Treat- ment Starts	3.95 3.90 2.30 2.55	4.5.5.0 0.00 0.00 0.00 0.00 0.00 0.00 0.	3.65 3.85 3.15	3.55 3.70 3.71 0.28
		Day 5	3.75 3.70 3.10 3.35	3.75 3.25 3.55	3.40 3.60 2.90 3.75	3.30 3.48 0.30
		Pay 4	44.6.6.0 0.40.4.	4	~	
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Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

APPENDIX N

INDIVIDUAL MATERNAL BODY WEIGHTS (kg) GROUP VII ORAL TECHNICAL ABATE

30 y		4.40	3,95	4.10	4.10	4.10		4.10	3.90		DEAD	8.8	4.10	4.35	4.11	0.16
Day 25		4.25	3,04	<u>ئ</u> س	4.04	4.11		4.70	3.95		3.62	4.01	3.45	4.22	4.02	0.18
23		4.25	3,94	3.99	6.0	4.05	DEAD	4.15	3.90		3.62	3.95	3.90	4.16	3.99	0.17
Day 19		4.05	3.92	3.90	3.86	4.01	2.12	4.03	3.84		3.38	3.57	3.80	4.08	3.71	Z
D& 18	Final Treat- ment	4.05	3.90	3.88	3.82	4.01	2.30	4.03	3.75		3.38	3.56	3.75	4.04	3.70	0.51
Day 17		4.05	3.87	3.88	3.8	4.01	2.32	4.03	3.66		3.36	3.56	3.70	4.04	3.69	0.48
Day 16		3.99	3.87	 8.	3.72	4.01	2.50	4.02	3.77		3.34	3.59	3.68	4.02	3.70	0.43
Day 15		3.97	3.84	3.96	3.77	3.96	2.60	3.90	3.94		3.38	3.45	3.63	3.90	3.69	0.40
Day 14		3.94	3.81	3.89	3.69	3.92	2.76	3.86	3.63		3.30	3.41	3.60	3.85	3,64	0.34
Z Day 13		3.88	3.80	3.89	3.66	3.95	2.17	3.71	3.73		3.33	3.46	3.52	3.84	3,63	0.33
GESTATION Day 12		3.85	3.78	3.77	3.63	3.94	2.70	3.72	3.68		3.31	3.30	3,48	3.86	3,59	0.35
Day 11		3.72	3.74	3.84	3.70	3.89	2.80	3.70	3.65		3.29	3.40	3.43	3.84	3.58	0.31
Day 10		3.72	May 3,74	3.80	3.70	3.99	2.80	3.65	3.75		3.31	3,35	3.40	3.86	3.59	0.32
ge 6		3.63	3.83	3.75	3,70	3.90	2.97	3.47	3.62		3.56	3.48	3.30	3.86	3.55	0.27
S &		3.63	2, 5, 2, 8, 2, 8,	3.75	3.70	3.90	2.97	3.47	3.62	Apr	3.26	3.48	3,30	3.86	3.56	0.28
Day 7		3.63	3.8	3.75	3.70	3.90	2.97	3.47	3.62	DEAD 30	3.26	3.48	3.30	3.86	3.55	0.27
Day 6	Treat- ment Starts	3.80	3.74	3.93	3.51	3.87	3.13	3.45	3.64	3.60	3.19	3.47	3.24	3.86	3.56	0.26
Day 5		3.70	3.7	3.90	3.55	3.83	3.25	3.47	3.60	3.87	3.20	3.54	3.63	3.92	3.62	0.23
Day 4		3.62	3.83	3.87	3.55	 81.8	3.28	3.52	3.60	3.32	3.20	3.52	3.77	۳. ع	3.63	0.24
	Anima] Number	88	8 8	96	ይ	86	. 88	83	901	101	102	103	104	105	1×	₽ +1

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

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Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

	90 %	3.92 3.92 3.92 3.92 3.92 4.03 3.92 5.31 5.31
	Day 25	2.53 3.53 3.54 3.57 3.57 3.57 3.57 4.08 4.08 3.79 0.43
	23 23	3.38 3.61 3.64 3.64 3.64 3.64 3.77 4.01 3.86 3.62 3.62
	19 Pay	2. 2. 2. 2. 2. 2. 2. 2. 2. 2. 2. 2. 2. 2
	Day 18 Final Treat- ment	25.2 2.33 2.54 2.54 2.54 2.55 2.55 3.55 3.55
	Day 17	644.000.000.000.000.000.000.000.000.000.
	Day 16	44.72.00.00.00.00.00.00.00.00.00.00.00.00.00
	Day 15	& & & & & & & & & & & & & & & & & & &
	Day 14	23.88.88.88.88.88.88.88.88.88.88.88.88.88
	Day 13	######################################
NOIL	Day 12	25.20 2.20 2.20 2.20 2.20 2.20 2.20 2.20
GESTATION	Day 11	
DAY OF	Dey 10	64 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8
	Day 9	
	Day 8	6.6.6.6.6.6.6.6.6.6.6.6.6.6.6.6.6.6.6.
	Day 7	6.53 4.53 6.54 6.55 6.55 6.55 6.55 6.55 6.55 6.55
	Day 6 Treat- ment Starts	3.372 3.372 3.372 3.333 303 3.333 3.333 3.333 3.333 3.333 3.333 3.333 3.333 3.333 3.333 3.
	Seg.	6.0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0
	Day 4	2.59 7.20 2.33 2.50 2.50 2.50 2.50 2.50 2.50 2.50 2.50
	Animel Number	122 123 124 127 128 129 130 131 131 132 133 134 135

APPENDIX P INDIVIDUAL MATERNAL BODY WEIGHTS (kg) GROUP IX DERMAL PYRAX WITH 2 PERCENT ABATE

APPENDIX Q RBC CHOLINESTERASE ACTIVITY
COMPARISON WITH TIME OF VARIOUS TREATMENTS WITH PYRAX CONTROL

C	Tuestment	Mean <u>d</u>		esterase Activ	
Group	Treatment			vith Pyrax Cont of Gestation	rui
		Day 6	Day 10	Day 19	Day 30 Sacrifice
	·	Pretest	Test Day 4	Test Day 13	Day
IV	Pyrax Control	7.52	8.40	8.12	8.69
	•	2.60	2.87	2.64	2.15
Y	Dermal Pyrax	7.57	8.35	4.68*	<u>5.79</u> *
	10% ABATE	2.15	2.24	1.89	1.44
		n = 28	n = 28	n = 26	n = 27
		t = 0.05	t = 0.06	t = 4.01	t = 4.30
VI	Dermal Technical	7.74	7.43	2.77*	2.40*
	ABATE	1.87	2.52	2.63	0.34
		n = 28	n = 27	n = 24	n = 20
		t = 0.27	t = 0.97	t = 5.18	t = 8.14
VII	Oral Technical	8.06	7.18	4.78*	5.66*
	ABATE	2.31	2.37	1.30	1.99
		n = 27	n = 25	n = 23	n = .22
		t = 0.59	t = 1.18	t = 3.97	t = 3.51
VIII	I.P. 6-AN	9.35	10.09	10.24*†	9.90
		2.38	2.43	2.33	2.31
		n = 28	n = 28	n = 25	n = 26
		t = 2.01	t = 1.74	t = 2.21	t = 1.44
IX	Dermal Pyrax	8.88	9.03	9.95	9.47
	2% ABATE	2.02	2.16	2.25	2.11
		n = 28	n = 28	n = 23	n = 24
		t = 1.60	t = 0.68	t = 1.85	t = 0.93

^{*} Significantly different from Pyrax control value significantlys at p<.05, as determined by student's "t" test. \dagger Cholinesterase activity value significantly higher than control.

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

INDIVIDUAL MATERNAL

APPENDIX R

RBC CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP IV DERMAL PYRAX CONTROL

	Day of Gestation						
Day 30	Day 19	Day 10	Day 6				
Sacrificed	Test Day 13 S	Test Day 4	Pretest	Animal Number			
7.8	8.5	11.7	8.4	46			
-	Dead	3.8	3.4	47			
9.6	10.3	9.5	9.0	48			
8.2	Missed	8.1	7.2	49			
11.0	8.4	8.7	7.4	50			
8.4	8.3	7.7	7.4	51			
9.2	7.7	8.8	6.9	52			
9.9	11.7	12.1	10.7	53			
10.2	9.5	9.8	8.9	54			
4.5	3.1	4.0	3.9	55			
12.8	11.8	12.9	12.7	56			
5.0	3.6	3.8	3.8	57			
7.5	6.9	7.3	6.4	58			
9.1	9.1	10.2	10.0	59			
7.3	6.7	7.6	6.7	60			
8.69	8.12	8.40	7.52	X			
2.15	2.64	2.87	2.60	+ SD			

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

INDIVIDUAL MATERNAL

APPENDIX S

RBC CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP V DERMAL PYRAX WITH 10 PERCENT ABATE

Day of Gestation						
	Day 6	Day	, 10	Day	19	Day 30
Pr	etest	Test	Day 4	Test	Day 13	Sacrificed
	8.3		8.8		3.2	3.9
	7.2		8.0		7.3	8.2
	9.3	•	10.4		4.8	7.2
	6.7	•	7.7		3.8	5.8
	9.6	•	10.2		4.0	5.3
	5.8	•	7.7		7.6	7.7
	3.1		3.9		3.0	4.4
	10.8		11.5		6.7	6.9
	9.7		0.5		5.0	5.5
	4.7	•	5.4		2.0	3.7
	9.0		9.4		7.5	7.6
	7.0		7.1		4.1	6.0
	5.9		6.4		4.8	5.0
	9.6	•	11.4		4.7	5.2
	6.8	•	6.8		1.7	4.4
	7.57		3.35	4	.68	5.79
	2.84		2.24		.89	1.44

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

INDIVIDUAL MATERNAL

APPENDIX T

RBC CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP VI DERMAL TECHNICAL ABATE

		Day of Ges		
Day 30	Day 19	Day 10	Day 6	
Sacrificed	Test Day 13	Test Day 4	Pretest	Animal Number
2.5	1.1	3.4	5.4	75
Dead	1.8	10.5	8.8	76
2.8	1.1	8.0	7.4	77
Dead	1.5	9.0	9.7	78
Dead	3.4	9.8	7.3	79
2.1	0.71	4.3	9.3	80
2.3	4.1	7.5	6.4	81
3.0	2.9	6.1	6.4	82
		Dead	6.2	83
2.2	10.1	10.1	9.7	84
2.2	1.5	8.5	9.6	85
Dead	0.82	3.9	6.3	86
	Dead	4.4	4.3	87
Dead	1.5	9.5	10.5	88
2.1	5.5	9.0	8.8	89
2.40	2.77	· 7.43	7.74	x
0.34	2.63	2.52	2.23	<u>±</u>

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

INDIVIDUAL MATERNAL

APPENDIX U

RBC CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP VII ORAL TECHNICAL ABATE

		Day of Ges	tation	
	Day 6	Day 10	Day 19	Day 30
Animal Number	Pretest	Test Day 4	Test Day 13	Sacrificed
91	7.7	7.9	5.6	5.8
92	9.1	DEAD	-	-
93	4.4	4.1	3.7	3.4
94	6.7	6.4	4.0	3.3
95	9.4	8.1	5.7	5.5
96	7.6	6.6	3.8	4.4
97	Dead	~	**	_
98	4.7	2.7	4.8	_
99	5.0	5.9	4.2	3.7
100	10.9	9.4	5.6	8.7
101	10.0	DEAD	-	-
102	7.1	5.8	4.0	-
103	9.0	9.7	3.8	6.1
104	11.9	9.0	4.0	7.4
105	9.4	10.6	8.1	8.3
x	8.06	7.18	4.78	· 5 . 66
+ SD	2.31	2.37	1.30	1.99

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

INDIVIDUAL MATERNAL

APPENDIX V

RBC CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP VIII INTRAPERITONEAL 6-AN

Day of Gestation				
Day 30	Day 19	Day 10	Day 6	
Sacrificed			Pretest	Animal
	Test Day 13	Test Day 4		Number
_	DEAD	12.2	13.3	106
10.3	10.4	10.7	10.2	107
10.7	11.7	11.8	10.6	108
8.4	9.1	8.6	7.4	109
8.0	7.9	8.5	7.4	110
14.9	14.6	15.4	12.4	111
10.2	12.0	10.8	9.7	112
9.7	9.1	8.3	7.8	113
9.7	9.6	8.6	7.6	114
9.8	10.7	9.9	10.1	115
14.7	14.5	13.9	13.4	116
8.7	10.0	10.0	9.7	117
7.8	7.1	7.2	6.4	118
8.2	9.5	8.8	8.3	119
7.5	7.2	6.7	5.9	120
9.90	10.24	10.09	9.35	ž
2.31	2.33	2.43	2.38	± SD

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

INDIVIDUAL MATERNAL

APPENDIX W

RBC CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP IX DERMAL PYRAX WITH 2 PERCENT ABATE

	tation	Day of Ges		
Day 30	Day 19	Day 10	Day 6	
Sacrificed	Test Day 13	Test Day 4	Pretest	Animal Number
10.2	12.2	10.2	9.7	121
10.0	11.7	9.6	10.5	122
6.9	7.1	5.1	5.8	123
8.0	7.4	6.0	5.7	124
7.1	8.4	8.0	8.0	125
7.7	7.5	7.1	6.8	126
11.9	10.9	9.7	10.1	127
7.9	7.4	6.9	6.8	128
	DEAD	9.6	9.8	129
14.0	12.8	12.6	11.0	130
9.6	10.4	8.9	10.2	131
9.5	10.8	11.1	10.6	132
-	DEAD	12.0	8.1	133
_	DEAD	8.1	7.8	134
10.8	12.8	10.5	12.3	135
9.47	9.95	9.03	8.88	x
2.11	2.25	2.16	2.02	± SD

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

APPENDIX X PLASMA CHOLINESTERASE ACTIVITY COMPARISON WITH TIME OF VARIOUS TREATMENT WITH PYRAX CONTROL

Group	Treatment	Mean (±SD) Plasma Cholinesterase Activity Values Comparison With Pyrax Control				
		D C	Day of Gest	cation Day 19	Day 30	
		Day 6 Pretest	Day 10 Test Day 4	Test Day 13	Sacrifice Day	
IV	Pyrax Control	4.97	3.34	3.75	2.71	
	•	1.18	0.81	1.01	1.08	
٧	Dermal Pyrax	5.01	3.07	2.52*	2.97	
	10% ABATE	1.01	1.06	$\overline{1.01}$	0.99	
		n = 28	n = 28	n = 26	n = 27	
		t = 0.08	t = 0.77	t = 3.24	t = 0.67	
٧I	Dermal Technical	5.56	2.97	1.34*	1.54*	
	ABATE	0.90	1.14	$\overline{0.90}$	0.63	
		n = 28	n = 27	n = 24	n = 20	
		t = 1.53	t = 1.01	t = 6.45	t = 2.80	
VII	Oral Technical	5.59	4.2 <u>7</u> *†	3.53	2.97	
	ABATE	0.91	0.99	0.90	1.08	
		n = 27	n = 25	n = 23	n = .22	
		t = 1.56	t = 2.68	t = 0.58	t = 0.57	
VIII	IP 6-AN	5.75	5.11*†	5.38*†	4.57*†	
		1.31	$\overline{1.35}$	1.34	1.19	
		n = 28	n = 28	n = 25	n = 26	
		t = 1.70	t = 4.36	t = 3.54	t = 4.32	
IX	Dermal Pyrax	5.56	4.56*†	4.88*†	2.75	
	2% ABATE	1.15	1.13	0.74	0.84	
		n = 28	n = 28	n = 23	n = 24	
		t = 1.38	t = 3.41	☆ ~ 3.16	t = 0.09	

^{*} Significantly different from Pyrax control values at p<.05 as determined by student's "t" test.
† Cholinesterase activity significantly higher than control.

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

INDIVIDUAL MATERNAL

APPENDIX Y

PLASMA CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP IV DERMAL PYRAX CONTROL

-	Day 6	Day o Day 10	f Gestation Day 19	Day 30
Animal	Pretest	Test Day 4	Test Day 13	Sacrificed
46	4.9	3.3	4.4	3.7
47	5.6	4.6	Dead	Dead
48	3.9	2.6	2.6	2.0
49	4.0	2.7	Missed	1.7
50	4.0	2.2	3.0	2.0
51	5.8	3.9	5.3	4.0
52	5.1	4.3	4.1	2.7
53	6.0	4.1	4.1	4.1
54	3.6	2.2	2.9	2.1
55	4.7	3.3	2.8	1.9
56	4.0	2.3	2.3	1.8
57	5.8	3.4	4.1	2.4
58	5.8	3.9	4.2	2.4
59	3.6	3.1	3.5	1.9
60	7.8	4.2	5.5	4.7
ž	4.97	3.34	3.75	2.71
+ SD	1.18	0.81	1.01	1.08

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INDIVIDUAL MATERNAL

PLASMA CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS)

GROUP V

DERMAL PYRAX WITH 10 PERCENT ABATE

		Day of Gestation							
	Day 6	Day 10	Day 19	Day 30					
Animal	Pretest	Test Day 4	Test Day 13	Sacrificed					
61	6.2	3.4	2.0	1.7					
62	5.0	2.4	2.9	2.2					
63	5.8	1.5	3.2	3.8					
64	4.0	2.5	1.5	2.5					
65	4.8	3.8	2.6	3.0					
66	3.9	2.8	2.8	2.2					
67	7.3	5.7	2.8	3.9					
68	4.8	3.5	2.9	2.5					
69	4.4	3.3	1.6	2.0					
70	4.8	3.3	1.8	3.4					
71	4.0	2.9	2.1	3.5					
72	6.3	3.4	3.5	4.3					
73	5.4	3.9	5.0	4.7					
74	4.1	2.5	2.3	3.5					
90	4.3	1.2	0.7	1.4					
x	5.01	3.07	2.52	2.97					
+ SD	1.01	1.06	1.01	0.99					

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INDIVIDUAL MATERNAL

PLASMA CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS)
GROUP VI
DERMAL TECHNICAL ABATE

Day of Gestation						
	Day 6	Day 10	Day 19	Day 30		
Animal	Pretest	Test Day 4	Test Day 13	Sacrificed		
75	5.2	1.2	1.0	1.4		
76	5.1	2.6	0.5	DEAD		
77	5.9	3.5	0.6	1.4		
78	5.6	2.1	0.7	DEAD		
79	5.7	3.1	2.5	DEAD		
80	5.0	1.2	0.5	1.3		
81	4.6	2.6	1.5	1.8		
82	6.3	3.1	1.3	3.0		
83	6.6	DEAD				
84	4.3	3.4	3.1	1.1		
85	7.0	3.9	1.1	DEAD		
86	6.6	2.5	1.7	1.2		
87	5.7	5.7	DEAD			
88	3.8	2.9	0.4	DEAD		
89	6.0	3.8	2.6	1.1		
x	5.56	2.97	1.34	1.54		
+ SD	0.90	1.14	0.90	0.63		

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INDIVIDUAL MATERNAL

APPENDIX BB

PLASMA CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP VII ORAL ABATE

	Day of Gestation						
	Day 6	Day 10	Day 19	Day 30			
Animal	Pretest	Test Day 4	Test Day 13	Sacrificed			
91	3.9	3.6	3.6	1.3			
32	5.6	DEAD	~ ^ -				
3	6.0	5.3	3.5	3.3			
94	5.7	5.2	4.0	2.0			
95	4.7	3.3	2.8	2.5			
96	6.6	4.7	3.0	3.4			
97	Dead	-	~	-			
98	5.5	2.4	1.9	De ad			
99	6.3	4.6	5.3	5.3			
100	4.6	3.5	3.4	3. 5			
101	4.5	DEAD	-	-			
102	6.4	4.6	3.9	-			
103	5.1	4.4	2.6	3.3			
104	6.6	3.7	3.9	2.6			
105	6.7	5.9	4.5	2.5			
į	5.59	4.27	3.53	2.97			
+ SD	0.91	0.99	0.90	1.08			

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INDIVIDUAL MATERNAL
PLASMA CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS)
GROUP VII
INTRAPERITONEAL 6-AN

	Day 6	Day o Day 10	f Gestation Day 19	Day 30	-
Animal	Pretest	Test Day 4	Test Day 13	Sacrificed	
106	4.7	4.6	Dead		
107	4.9	3.9	3.9	3.0	
108	9.7	9.6	9.1	7.6	
109	6.7	5.5	5.5	5.5	
110	5.0	4.1	4.3	3.8	
111	6.5	4.8	7.3	4.5	
112	5.3	4.2	5.2	5.5	
113	5.2	4.7	5.1	3.9	
114	6.1	4.8	4.9	3.7	
115	5.0	4.6	5.3	5.3	
116	5.3	5.7	5.5	4.7	
117	4.4	4.9	4.7	4.7	
118	5.8	4.8	4.6	3.9	
119	6.6	5.8	5.5	4.8	
120	5.0	4.7	4.4	3.1	
x	5.75	5.11	5.38	4.57	
<u>+</u> SD	1.31	1.35	. 1.34	1.19	

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INDIVIDUAL MATERNAL

APPENDIX DD

PLASMA CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS) GROUP IX DERMAL PYRAX WITH 2 PERCENT ABATE

	Day 6	Day o	f Gestation Day 19	Day 30	
	bay 0	Day 10	Day 19	Day 30	
Animal	Pretest	Test Day 4	Test Day 13	Sacrificed	
121	4.5	3.6	4.0	1.4	
122	5.7	4.5	5.0	2.7	
123	6.4	5.5	5.0	2.4	
124	6.5	4.5	4.9	2.5	
125	4.8	3.4	5.8	4.5	
126	5.8	5.5	6.0	2.8	
127	6.9	4.9	4.8	3.1	
128	5.2	4.8	3.8	3.5	
129	5.6	4.3	DEAD	_	
130	7.0	5.4	5.3	2.4	
131	6.9	5.7	5.6	2.3	
132	5.2	4.6	4.4	1.8	
133	2.9	1.6	DEAD	=	
134	5.9	6.1	DEAD	_	
135	4.1	4.0	3.9	3.6	
x	5.56	4.56	4.88	2.75	
+ SD	1.15	1.13	0.74	0.84	

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MATERNAL BRAIN CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS)
AT TERMINATION OF PREGNANCY (Day 30 of Gestation)

APPENDIX EE

Group IX Dermal Pyrax 2% ABATE (0.81 gm/kg/day)	Activity	49.2 57.3 54.3 54.3 52.9 52.9
Gro De- Py 2% 19.	Animal	121 122 123 124 126 126 130 131 132 135
Group VIII Positive Control IP 6-AN	Activity	53.0 47.0 48.7 48.7 50.7 55.3 55.3 53.7 51.3
Groun Pos Con IP	Animal Number	107 108 109 110 111 1112 1113 1114 1116 117 118
Group VII Oral ABATE 2 mg/kg/day)	Activity	47. 6 49. 0 52. 6 47. 4 45. 3 65. 4 65. 4
Grou 0 AB (32 mg	Animal Number	96 96 96 100 104 105 105
Group VI Dermal Technical ABATE (0.14 ml/kg/day)	Activity	39.3 40.6 42.0 38.7 37.6 41.3
Gro De Tech AB (0.14 m	Animal Number	75 77 77 80 81 84 86 89
Group V Dermal Pyrax 10% ABATE 81 gm/kg/day)	Activity	8.8.8.4.4.4.6.9.9.9.9.9.9.9.9.9.9.9.9.9.9.9.9
Group V Dermal Pyrax 10% ABAT (1.81 gm/kg	Animal Number	65 65 65 65 65 65 65 72 73 73 73
Group IV Dermal Pyrax Control (1.81 gm/kg/day)	Activity	46.3 49.0 47.3 48.1 48.9 50.6 55.7
Group Dermal Pyrax Control	Animal Number	55 55 55 55 56 56 56 56 56 56 56 56 56 5
•		

APPENDIX FF

MATERNAL BRAIN CHOLINESTERASE ACTIVITY (GARRY AND ROUTH UNITS)

COMPARISON WITH TIME OF VARIOUS TREATMENTS WITH

PYRAX CONTROL

Group	Treatment	Mean &SD) Brain Cholinesterase Activity Values at End of Study (Day 30 of Pregnancy)			
			DF	t	
Group IV	Dermal Pyrax Control	48.7 2.9	-	••	
Group V	Dermal Pyrax 10% ABATE	51.2 4.4	26	1.75	
Group VI	Dermal Technical ABATE	38.9 3.5	19	6.89*	
Group VII	Oral ABATE	49.0 2.9	21	0.24	
Group VIII	IP 6-AN	51.3 3.5	25	3.09*†	
Group IX	Dermal Pyrax 2% ABATE	51.5 3.0	23	2.31*†	

 $[\]star$ Significantly different from Pyrax control values at p<.05 as determined by students "t" test.

t Cholinesterase activity values higher than control.

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APPENDIX GG

MEAN FETAL BODY WEIGHT AND LENGTH PER DOE

	Animal		t (gm)	Length (cm)	
Group	Number	X	± SD	<u> </u>	±SD
17	46	_	_	_	-
Dermal	48	44.4	6.6	8.5	0.5
(Control Pyrax)	49	42.2	4.5	8.5	0.3
(conc. or igian)	50	55.4	3.8	8.2	0.1
	51	-	-	-	-
	52	40.4	3.6	8.7	0.5
	53	-	-	-	-
	54	46.3	12.9	8.5	0.5
	55	63.1	3.1	9.1	0.6
	56	44.9	8.9	8.9	0.5
	57	57.0	3.9	9.3	0.6
	57 58	50.9	5.7	9.3 9.4	0.8
	59	51.9	8.2	8.8	
		31.9			0.4
	60	-	-	-	
٧	61	37.8	6.5	8.1	0.7
Dermal Pyrax	62	58.0	4.6	8.9	0.2
10% ABATE	63	-	_	-	-
	64	45.4	8.3	8.4	0.7
	65	53.8	10.4	9.0	0.8
	66	53.9	5.1	8.8	0.7
	67	49.2	10.5	8.9	0.7
	70		-	-	-
	71	_	-	~	•
	72	54.8	5.6	9.6	0.5
	73	-	-	-	-
	74		_	_	
	90	48.9	12.0	8.5	0.7
	30	40.5	12.0	0.5	0.7
V I	75	33.2	6.4	8.7	0.8
Dermal Technical	76	-	•	-	- -
ABATE	77	46.1	15.2	9.9	1.6
	81	-		_	_
	82	-	-	•=	-
	84	41.6	4.5	8.4	0.4
	86	36.8	3.5	8.2	0.3
	89	46.1	7.2	8.4	0.4
	• • • • • • • • • • • • • • • • • • • •	10.2	,	V• T	0. 1

Phase 4, Toxicological Assessment of ABATE, Study No. 75-51-1302-84, Apr 83

	Animal	Weigh	t (gm)	Length (cm)	
Group	Number	<u> </u>	<u>+</u> SD	X	<u>±</u> SD
VII	91	53.6	3.5	9.1	0.5
Oral ABATE	93	40.4	6.6	8.7	0.7
(32 mg/kg/day)	94	54.3	4.7	9.3	0.4
(029, 1.9, 0.0)	95	49.5	8.8	9.4	0.6
•	96	47.4	5.1	9.1	0.4
	99	-	-	-	-
	100	-	_	-	_
	103	46.2	4.6	9.1	0.7
	104	46.4	4.9	9.1	0.3
	105	59.8	9.1	9.7	0.5
VIII	107	44.1	3.7	8.2	0.8
IP 6-AN	108	77.1	J. /	O• <i>t</i>	0. 0
IF U-AN	109	_		_	_
	110	_	-	-	_
	111	_			-
	112	_	<u></u>	_	
	113	36.1	3.3	7.5	0.3
	114	21.6	5.2	6.6	0.6
	115	_	-	_	-
	116	_	-	-	-
	117	-	-	-	~
	118	35.1	14.3	6.9	1.3
	119		-	_	•
	120	41.5	. 5.3	7.1	1.5
IX	121	51.0	6.2	9.2	0.8
Dermal Pyrax	122	61.8	3.9	9.6	0.6
2% ABATE	123	61.7	4.3	9.4	0.6
	124	49.9	4.3	9.8	0.6
	125	-	-		-
	126	49.0	5.2	8.9	0.7
	127	53.5	5.8	9.4	0.9
	128	44.9	4.9	8.6	8.0
	130	54.0	3.4	9.6	0.5
	131	55. 5	5.9	9.6	0.6
	132	48.6	9. 8	9.0	0.7
	135	-	~	-	••

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APPENDIX HH

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